REVIEW ARTICLE



WILEY

Treatment strategy for isolated bile leakage after hepatectomy: Literature review

Norio Kubo 💿 | Ken Shirabe

Department of Hepatobiliary and Pancreatic Surgery, Gunma University Graduate School of Medicine, Maebashi, Japan

Correspondence

Ken Shirabe, Department of Hepatobiliary and Pancreatic Surgery, Gunma University Graduate School of Medicine, 3-39-22 Showa-machi, Maebashi, Gunma 371-8511, Japan. Email: kshirabe@gunma-u.ac.jp

Abstract

Isolated bile leakage (IBL) after hepatectomy is intractable, and various treatment methods for it have been reported. This review aimed to clarify the treatment strategy for IBL by summarizing studies on IBL after hepatectomy without extrahepatic bile duct resection. Thirty-three cases of IBL were reported. The incidence of IBL is very low, accounting for 0.1%-1% of all hepatectomy cases. The risk factors for IBL are unclear; however, several reports mention that biliary anomaly is associated with a high risk of IBL, with preoperative and intraoperative confirmation of biliary tree anatomy being the most important preventive strategy. Treatment methods for IBL include liver resection, bilioenteric anastomosis, endoscopic treatment, bile duct ablation, percutaneous transhepatic portal vein embolization (PTPE), transcatheter arterial embolization, and use of fibrin glue. The therapeutic methods should be chosen depending on remnant liver function, amount of bile leakage, and the liver volume causing the bile leakage. When there is bile leakage from less than one segment, nonsurgical treatment is recommended, whereas when there is bile leakage from one or more segments, surgical treatment can be recommended. Nevertheless, recently, non-surgical treatment such as PTPE, PTPE with bile duct ablation, and endoscopic methods have been considered as effective treatment approaches.

KEYWORDS

anatomical variation, bile leakage, divided bile ducts, hepatectomy, isolated bile duct

1 | INTRODUCTION

Currently, hepatectomy can be carried out safely because of the improvement in surgical techniques and perioperative management.^{1,2} Mortality rates after hepatectomy have decreased from 2.6% to 1.6%.^{1,3} However, the incidence of bile leakage has been reported to be 3.3%–8.7% and has not changed over the past few decades.¹⁻⁵ According to the International Study Group of Liver Surgery (ISGLS), bile leakage after hepatectomy is defined as drainage of fluid with bil-irubin level three times greater than the serum level at postoperative

day 3 or the need for interventions owing to bilious collection or biliary peritonitis.⁶ Postoperative bile leakage is associated with an increased risk of post-hepatectomy liver failure and mortality.^{7,8}

Most bile leakage cases are treated with simple drainage. However, some types of bile leakage require interventions, such as endoscopic bile drainage or percutaneous abdominal drainage. Intractable bile leakage is defined as bile leakage that does not improve after a drainage procedure. Bile leakage was classified by Nagano et al into the following four groups: type A, minor leakage from a cut surface; type B, leakage caused by insufficient closure of the bile duct stump; type C, leakage from the injured bile duct wall at

This is an open access article under the terms of the Creative Commons Attribution-NonCommercial-NoDerivs License, which permits use and distribution in any medium, provided the original work is properly cited, the use is non-commercial and no modifications or adaptations are made. © 2019 The Authors. Annals of Gastroenterological Surgery published by John Wiley & Sons Australia, Ltd on behalf of The Japanese Society of Gastroenterology

-WILEY- AGSurg Annals of Gastroenterological Surgery

the exposed bile duct or hilar bile duct; and type D, leakage from the distal orifice of the isolated bile duct.^{4,9} Type D bile leakage is intractable and cannot be treated by simple drainage alone. Patients with Type D bile leakage undergo surgical procedures, such as liver resection or choledochojejunostomy, or non-surgical procedures, such as bile duct ablation with absolute ethanol, transcatheter arterial embolization (TAE), or percutaneous transhepatic portal embolization (PTPE) of the liver segment that produced the bile leak. We summarized the reports of leakage from the isolated bile duct (isolated bile leakage [IBL]), regardless of the size of the independent bile duct, after hepatectomy to establish treatment strategies for IBL.

2 | LITERATURE SEARCH

PubMed (URL: https://www.ncbi.nlm.nih.gov/pubmed) was searched for previous reports on IBL after hepatectomy that were published from 1998 to June 2018. The search was carried out using the terms "bile leakage" and "hepatectomy." Through the literature search, no systematic reviews on IBL were found, and there were 29 studies reporting IBL, including 33 cases of IBL. We focused on the management of IBL. These findings are summarized in Figure 1.

3 | RISK FACTORS FOR IBL

Risk factors for bile leakage have been described in several studies.^{4,5} IBL was very rare and accounted for 0.1%–1% of all hepatectomy cases in some reports.^{10–13} Risk factors for IBL have not been reported clearly. Types of liver resection in patients with IBL were various, including 22 cases of major hepatectomy and five cases of minor hepatectomy (Table 1). Bile duct anomalies were reported as a risk factor for IBL.^{11–15} After right-side major hepatectomy, IBL occurred from the bile duct of segment 1, left segmental bile duct, and remnant right-side liver.^{10,12,15} Some cases in which Spiegel's lobe was joined to the right posterior section or the common bile duct were reported.^{13,15,16} Patrono et al reported a biliary anomaly of a connection of the segment 2 duct to the right bile duct.¹¹ After left-side major hepatectomy, IBL occurred from the posterior bile duct because of the damaged right posterior segment bile duct draining into the left duct.^{11,13,14} Several types of bile duct anomalies were related to IBL.

4 | PREVENTION OF IBL

It is most important to confirm preoperative biliary tree images and perioperative assessment to prevent IBL.¹⁷ Gadoliniumethoxybenzyl-diethylenetriamine penta-acetic acid (Gd-EOB-DTPA)-enhanced magnetic resonance (MR) imaging could be used for preoperative evaluation of bile duct anatomy in addition to conventional information on focal hepatic lesions.¹⁸ Moreover, intraoperative cholangiography through the cystic duct before ligation of the bile duct is very important to prevent bile duct injury such as posterior segment bile duct drain into the left duct. In several reports, residual segments after hepatectomy may have caused isolated bile duct leakage.^{10,19} Anatomical resection with meticulous surgical technique is important to prevent IBL such as that from the posterior bile duct after anterior resection.

5 | DIAGNOSTIC METHOD FOR IBL

It is very important to classify the type of bile leakage because the treatment method for each type differs.⁹ For the detection of biliary leakage, cross-sectional imaging studies, including ultrasound, computed tomography (CT), and magnetic resonance cholangiography, are used.^{20,21} Endoscopic retrograde cholangiopancreatography (ERCP),²² drip-infusion cholangiography with computed tomography (DIC-CT),²³ and Gd-EOB-DTPA-enhanced MR cholangiography²⁴ are very important for detecting the bile leakage point. Fistulograms are also helpful for determining the type of biliary leakage and the degree to which leakage can be controlled by drainage.⁹ Direct enhancement of the bile duct by ERCP is needed for the diagnosis of IBL to confirm the absence of a connection to the central bile duct.²⁴ Area of the independent bile duct was confirmed by these images. The liver volume causing the bile leakage was measured by CT after injecting the contrast medium. Gd-EOB-DTPA-enhanced MR cholangiography using delayed-phase images was effective for detecting the presence and location of active bile leaks.²⁵ In two recently published studies investigating the value of 20-30 minutes delayed Gd-EOB-DTPA-enhanced MR cholangiography for the detection of biliary complications after hepatobiliary surgery, both groups reported 100% sensitivity of this technique for the diagnosis of bile leakage.24,26

5.1 | Surgical treatment of IBL

The 11 cases that underwent surgical treatment for IBL are summarized in Table 2. Timing of surgical intervention was based on non-responsiveness to external drainage and/or the persistence of intra-abdominal sepsis.¹³ Surgical procedures were mostly carried out at several months after the first operation. Only one patient underwent the second operation at an early time point after the first operation.¹¹ Before the surgical treatment, percutaneous drainage, PTPE, and TAE were done to treat the IBL. A planned approach was based on the patient's general status, volume of future liver remnant and liver functional reserve, type and extent of injury, and volume of the causing IBL.¹³ A previous report mentioned that the decision for reoperation should be made as early as possible, preferably before the development of severe intra-abdominal sepsis and dense adhesions.⁸ It was difficult to immediately determine whether the IBL was not cured by non-surgical treatment. Surgical procedures were carried out in seven cases of liver resection of the independent liver parenchyma containing the fistula and in four cases of bilioenteric anastomosis.^{10-13,15,27} Another patient underwent resection with biliary-enteric anastomosis.^{10,13} Liver resection was done when the FIGURE 1 Extraction algorithm for the selection of articles reporting on isolated bile leakage

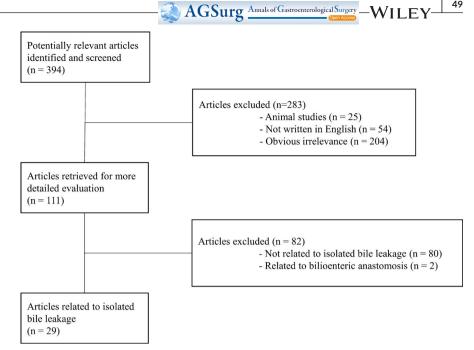


TABLE 1 Types of liver resection in patients with isolated bile leakage

Surgery	No. of isolated bile leakage cases	Leaked bile duct: number
Major		
Right extended hepatectomy	8	Left bile duct: 2, S1: 2. S2: 1. Remnant S5/8: 2, remnant S6: 1
Right extended hepatectomy	1	S1: 1
Right trisectionectomy	1	S2: 1
Left hepatectomy	3	Posterior: 1, S1: 1, remnant S4: 1
Left extended hepatectomy	2	S1/5: 1, posterior: 1
Left trisectionectomy	1	Posterior: 1
Central bisectionectomy	2	Posterior: 1, S8: 1
Right anterior sectionectomy	3	Posterior: 3
Right posterior sectionectomy	1	Anterior: 1
Minor		
Segmentectomy	3	Anterior: 1, Posterior: 1, ND: 1
Left lateral sectionectomy	1	Posterior + S4: 1
Wedge resection	2	S5: 1, ND: 1
Details of hepatectomy were unclear	5	S1: 1, ND: 4

Note: ND, not described.

independent liver parenchyma of the bile leakage was segment 1 or segment 4, and there was only one case in segments 6 and 7.

Bilioenteric anastomosis or fistulojejunostomy for IBL from the major bile duct such as posterior segment after left hemihepatectomy was reported in four cases in three reports.^{11,12,27} A percutaneous transhepatic drain was positioned into the excluded bile duct before operation, with the principal aim of guiding hilar plate dissection and facilitating the location of the excluded duct.¹¹ Anastomosis between the jejunum and the fistula was created using the drainage catheter as a guide.²⁷ Patrono et al reported that hepatico-jejunostomy on the

excluded bile duct presented the advantage of sparing unnecessary sacrifice of the liver parenchyma origin of the fistula.¹¹ When the flow of the bile duct is insufficient, patients undergoing bilioenteric anastomosis may be at risk for severe cholangitis and liver abscess after the operation.

Type of surgical procedure was based on the estimated volume of the liver remnant and the functional reserve of the liver and intraoperative factors, such as adhesions, infection, abscess formation, and anatomical distortions caused by regeneration of the remaining liver and anatomical errors of the first operation.^{10,13}

Complication

(p) 15

Postoperative hospital stay

Blood loss (mL)

Operating time (min)

Second operation Resection of S1

between 1st and 2nd operations

treatment Independent liver before 2nd

Period

Non-surgical

оZ

QN

QN

48 d

Percutaneous

Right hepatectomy S1 First operation

drainage

operation

segment

Diagnosis НСС

QN

Fukuhisa et al¹⁵

Rate of isolated bile leakage

> Reported year 2017

> > Author

Fragulidis et al ¹³	2008	3/234 (1%)	CCC	Right extended hepatectomy	S1	Percutaneous drainage	ę m	Resection of S1	QN	Q	DN	Uneventful
			Hydatid cyst	Resection of seg- ment 5	Posterior	Percutaneous drainage	8 m	Resection of S6 and S7	QN	QN	ND	Uneventful
			Hydatid cyst	Left lateral sectionectomy	S4, Posterior	Percutaneous drainage	14 m	Resection of S4 and biliary-enteric anastomosis	QN	QN	QN	Uneventful
Honore et al ¹⁰	2009	3/2409 (0.1%)	Hepatic abscess after laparoscopic cholecystectomy	Right hepatectomy	Remnant S 5/8	Percutaneous drainage	18 m	Resection of S5/8 and hepaticojejunostomy	401 4	450	30	Fistula on the bilio - digestive anastomosis
			HCC	Right hepatectomy	Remnant S 5/8	Percutaneous drainage	3 m	Resection of S5/8	310 2	2020	10	Uneventful
			HCC	Right hepatectomy	S6	PTPE, TAE and direct closure	12 m	Resection of S6	405 2	2300	13	Uneventful
Patrono et al ¹¹	2014	1	Hepatic injury	Left hepatectomy	S6/7	ENBD PTCD	Early timing	Bilioenteric anastomosis	QN	QN	DN	Uneventful
			Donor of LDLT	Right trisectionectomy	S2	PTCD	5 months	Bilioenteric anastomosis	QN	Q	ŊŊ	Uneventful
Hoekstra et al ¹²	2012	1/315 (0.3%)	Focal nodular hyperplasia	Right hepatectomy	Left segmental bile duct	I	I	Bilioenteric anastomosis	DN	QN	DN	Bile leakage
Sakamoto et al ²⁷	2016	2/334 (0.6%)	I	Right anterior sectionectomy	Posterior	Percutaneous drainage and PTPE	I	Fistulojejunostomy	QN	QN	323	DN
Abbreviati PTCD, perc	ons: CCC, cl cutaneous ti	holangiocellular ranshepatic cho	carcinoma; d, days; vlangio-drainage; PT	ENBD, endoscopic PE, percutaneous ti	nasobiliary drain. ranshepatic porta	age; HCC, hepato Il vein embolizatio	cellular carcir n; S, segment	Abbreviations: CCC, cholangiocellular carcinoma; d, days; ENBD, endoscopic nasobiliary drainage; HCC, hepatocellular carcinoma; LDLT, living-donor liver transplantation; m, months; ND, not described; PTCD, percutaneous transhepatic cholangio-drainage; PTPE, percutaneous transhepatic portal vein embolization; S, segment; TAE, transcatheter arterial embolization.	ver transplantat ial embolization	tion; m, ŀ.	months; ND, r	ot described;

TABLE 2 Surgical treatment for isolated bile leakage

5.2 | Non-surgical treatment of IBL

Non-surgical procedures to manage IBL are bile duct ablation, percutaneous PTPE or TAE of the liver segment that produced the bile leak. Non-surgical cases of IBL are summarized in Table 3.

5.2.1 | Endoscopic treatment

Generally, endoscopic drainage to the common bile duct is ineffective for IBL. Mutignani et al reported bridging stent treatment for IBL.²⁸ A transpapillary stent was inserted into the peritoneal cavity to drain the associated bilious collection, and a second stent was inserted into the bile duct to ensure proper biliary drainage for the rest of the liver.²⁸ Lee et al reported that fluoroscopy-guided transgastric hepaticoantrostomy was carried out for IBL after left hepatic trisectionectomy, and all external drainage catheters were removed.²⁹ The rendezvous procedure, which combines endoscopic techniques with percutaneous techniques for the treatment of IBL, was reported in only one study.³⁰ Biliary continuity was successfully restored in two out of the three patients using the rendezvous procedure.⁴⁰ These endoscopic treatments for IBL are minimally invasive procedures and cause little damage to liver function. However, there are only a few reports of endoscopic treatment because it is not widely carried out.

5.2.2 | Bile duct ablation therapy

Bile duct ablation therapy for IBL was mentioned in 13 case reports. Ethanol was commonly used for bile duct ablation therapy;^{30,31} acetic acid was used in one case and N-butyl cyanoacrylate in another.^{14,34} Kyokane et al reported that selective intrahepatic biliary ethanol injection destroyed the biliary epithelium, permeated the parenchyma, induced hepatocyte degeneration, and resulted in compensatory hypertrophy of the non-injective hepatic lobe in an animal study.⁴¹ Bile duct ablation therapy should only be done for IBL with no communication with the biliary tree because ethanol results in irreversible damage to the remaining bile ducts.^{30,32} It is necessary to confirm that the leaking bile ducts do not communicate with the biliary tree by carrying out fistulography and ERCP.²⁷ To prevent the outer bile duct from being exposed to ethanol, some authors used a balloon occlusion catheter.^{8,14,33} A previous study reported that ethanol injection into the liver parenchyma by a percutaneous transhepatic approach, instead of into the bile duct, destroyed both the biliary epithelium and liver parenchyma.35

Most cases of IBL that were treated with ethanol ablation therapy involved less than one segment. The largest area of IBL treated with ethanol ablation was the anterior or posterior segment.^{14,27,32,36} Sadakari et al reported that bile duct ablation with ethanol to the posterior segment was ineffective.³⁶ They carried out PTPE in the posterior portal branch. Shimizu et al treated IBL from the anterior bile duct with 23 attempts of bile duct ablation AGSurg Annals of Gastroenterological Surgery -WILEY

with ethanol.³² Sakamoto et al reported that they successfully treated a case with posterior bile duct ablation with ethanol as the treatment process gradually decreased the size of the residual posterior segment.²⁷ When the liver volume is large, or when the amount of leaked bile is high, IBL may often not be cured by ethanol ablation therapy.

5.2.3 | Percutaneous transhepatic portal vein embolization

Percutaneous transhepatic portal vein embolization induces atrophy of hepatocytes and decreases the amount of bile duct juice.³⁶ PTPE was reported in five cases; of these, three cases were successfully treated,^{23,36,37} whereas the other two cases underwent operation without achieving successful treatment of the bile leakage.^{10,27} PTPE was suitable for IBL from one or more liver segments. PTPE with fibrin glue or ethanol was reported to treat isolated bile duct leakage from the posterior segment.^{36,37} Thus, PTPE was used to decrease the amount of bile leakage from a large area of the isolated bile duct when ethanol injection was ineffective.^{10,27,37} However, some reports have shown that bile leakage was not stopped after PTPE and needed to be treated with surgery.^{10,27} In a previous study, treatment with ethanol injection to the fistula before PTPE failed because of the large amount of bile leakage. Combination therapy with bile duct ablation after PTPE was reported to be useful.²³ PTPE should be done only when patients are in good condition, with sufficient remnant liver function to avoid liver failure.

5.2.4 | Transcatheter arterial embolization

Treatment for IBL with TAE was mentioned in only one successfully treated case and in an unsuccessfully treated case.^{10,38} TAE in the anterior segmental artery was carried out to stop the production of bile in the injured part of the anterior segment after simple drainage and ethanol injection treatment failed.³⁸ The TAE treatment was effective, and the patient was discharged 15 days after TAE. In contrast, Honore et al reported a case in which liver resection was carried out as the definitive treatment because TAE 1 month after PTPE failed to stop bile leakage.¹⁰ Patients with a liver abscess after TAE are at high risk of developing bile duct infection because of liver parenchyma necrosis after TAE.⁴² When bile leakage was detected after hepatectomy, almost all cases had abdominal infections. TAE may be adaptable only when the bile leakage area is very small; otherwise, infectious complications may arise.

5.2.5 | Fibrin glue

Tanaka et al reported two cases of fibrin glue treatment for IBL.³⁹ After confirmation that the fistula was free of infection and that

Author	Reported year	Rate of isolated bile leakage	Diagnosis	First operation	Independent liver segment	Treatment method	Outcome
Kyokane et al ³⁰	2002	DN	Gallbladder carcinoma	Right hepatectomy	S2	Ethanol injection	
Sakaguchi et al ³¹	2011	I	Liver metastasis from GIST	Extended left hepatectomy	S5 + 1	Ethanol injection	
Shimizu et al ³²	2006	I	HCC	Right posterior sectionectomy	Anterior bile duct	Ethanol injection	Alive
Matsumoto et al ³³	2002	I	HCC	Right hepatectomy	Caudate lobe	Ethanol injection	Alive
Nakagawa et al ⁴	2017	1/631 (0.2%)	1	I	I	Ethanol injection	
Kusano et al ¹⁹	2003	I	Liver abscess with intrahe- patic stones	Left hepatectomy	S4	Ethanol injection	Alive
Yamashita et al ⁸	2001	3/781 (0.4%)	1	I	1	Ethanol injection with balloon catheter occlusion	
			1	I	I	Ethanol injection	
			I	I	I	Ethanol injection	
Sakamoto et al ²⁷	2016	2/334 (0.6%)	I	Right anterior sectionectomy	Posterior	Ethanol injection	Alive
Park et al ¹⁴	2005	I	Biliary cystadenocarcinoma	Left extended hepatectomy	Posterior	Acetic acid	Alive
Kim et al ³⁴	2012	I	HCC	Central bisectionectomy	S8	N-butyl cyanoacrylate	Alive
Kataoka et al ³⁵	2011	I	HCC	S5 segmentectomy	I	Ethanol injection into the liver parenchyma	Alive
Kubo et al ²³	2018	1	HCC	Partial hepatectomy of S4/5	S5 + 8	Combination therapy with etha- nol injection and PTPE	Alive
Sadakari et al ³⁶	2008	I	Liver metastasis from rectal cancer	Central bisectionectomy	Posterior	ртре	Alive
Hai et al ³⁷	2012	I	HCC	Right anterior sectionectomy	Posterior	PTPE	Alive
lkeda et al ³⁸	2015	I	Gallbladder cancer	Extended cholecystectomy	S5	TAE	
Tanaka et al ³⁹	2002	2/363 (0.6%)	CCC	Left hepatectomy	Caudate branch	Fibrin glue	Alive
			HCC	Partial hepatectomy	I	Fibrin glue	Alive
Mutignani et al ²⁸	2017	I	Cholangiocarcinoma	Right hepatectomy	Left lobe branch	Bridging stent	
Lee et al ²⁹	2015	I	НСС	Left trisectionectomy	Posterior	Fluoroscopy-guided transgastric hepaticoantrostomy	

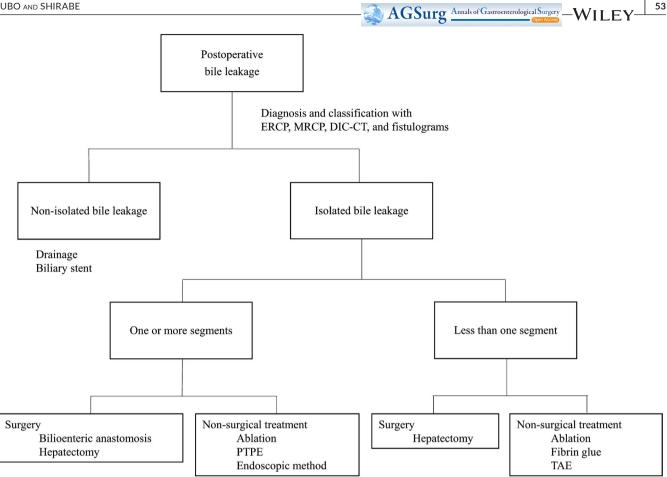


FIGURE 2 Postoperative bile leakage was diagnosed and classified by endoscopic retrograde cholangiopancreatography (ERCP), magnetic resonance cholangiopancreatography (MRCP), drip-infusion cholangiography with computed tomography (DIC-CT), and fistulograms. Therapeutic strategy of isolated bile leakage was classified by the quantity of bile leakage and as either surgical or non-surgical treatment. When the isolated bile leakage was from less than one liver segment, the first choice of treatment method was non-surgical treatment such as ethanol ablation. When the isolated bile leakage was from more than one segment, the first choice of treatment method was surgical treatment. PTPE, percutaneous transhepatic portal vein embolization; TAE, transcatheter arterial embolization

the volume was less than 50 mL/day, the fistula was completely sealed with a mixture of fibrin glue and iodized oil (Lipiodol; Kodama Pharmaceutical, Tokyo, Japan). The fistula was immediately closed without any major complications. It was considered that patients were required to have a low volume of bile leakage for treatment with fibrin glue. Treatment with fibrin glue was reported in only two cases, and the treatment effect was limited.

TREATMENT STRATEGY FOR IBL 6

We summarized the treatment strategy for IBL used in previous reports (Figure 2). IBL was confirmed by the absence of continuity. When the bile duct was connected, drainage and biliary stenting were carried out to control bile leakage. However, how do we choose the optimal treatment method for cases with unconnected bile duct? The most effective and minimally invasive treatment method should be chosen depending on the liver volume causing the bile leakage and liver function. When the liver volume causing bile leakage, as assessed using DIC-CT,

fistulograms, and ERCP, is less than one segment, non-surgical treatment with bile duct ablation should be initially considered. Fibrin glue or TAE should be restricted for cases with IBL from a very small region. Some cases reported successful treatment of IBL by liver resection.^{8,10,13,15}

When the liver volume causing the bile leakage is one or more segments, fewer cases are amenable to treatment with bile duct ablation or combination treatment with bile duct ablation and PTPE.²³ If the non-surgical method is ineffective, surgical procedures, such as bilioenteric anastomosis, fistulojejunostomy, or liver resection should be carried out. However, adhesions as a result of inflammation make these operations difficult, especially in cases with cirrhosis. Therefore, the operation should be done by experienced surgeons.

One limitation of the present study is that since we only summarized the reports that were available on PubMed, almost all these cases were successfully treated. Thus, there may be publication bias against cases wherein no treatment modality was successful. Further study will be needed to confirm the optimal strategy for the treatment of IBL.

7 | CONCLUSIONS

Isolated bile leakage is a very rare complication. This type of bile leakage is intractable and results in a longer hospital stay. It is important that patients with IBL be diagnosed definitively and early in order to give appropriate treatment. Moreover, it is necessary to be careful in cases of bile duct anomalies such as posterior segment bile duct drainage into the left duct. Intraoperative bile duct enhancement to confirm the biliary tree before resection of the bile duct is very important to prevent IBL. The treatment method should be chosen depending on the liver volume causing the bile leakage. When the IBL is from less than one liver segment, non-surgical treatment may be recommended as the first choice. Conversely, when the IBL is from one or more segments, in some cases, IBL may be treated by non-surgical treatment such as PTPE or bile duct ablation. If the non-surgical method is ineffective, surgical procedures, such as bilioenteric anastomosis, fistulojejunostomy, or liver resection, should be carried out.

-WILEY- AGSurg Annals of Gastroenterological Surgery

DISCLOSURE

Conflicts of Interest: Authors declare no conflicts of interest for this article.

ORCID

Norio Kubo (D) https://orcid.org/0000-0003-2624-2529

REFERENCES

- Brooke-Smith M, Figueras J, Ullah S, Rees M, Vauthey J-N, Hugh TJ, et al. Prospective evaluation of the International Study Group for Liver Surgery definition of bile leak after a liver resection and the role of routine operative drainage: an international multicentre study. HPB (Oxford). 2015;17:46–51.
- Kyoden Y, Imamura H, Sano K, Beck Y, Sugawara Y, Kokudo N, et al. Value of prophylactic abdominal drainage in 1269 consecutive cases of elective liver resection. J Hepatobiliary Pancreat Sci. 2010;17:186–92.
- Yang T, Zhang J, Lu JH, Yang GS, Wu MC, Yu WF. Risk factors influencing postoperative outcomes of major hepatic resection of hepatocellular carcinoma for patients with underlying liver diseases. World J Surg. 2011;35:2073–82.
- Nakagawa K, Tanaka K, Nojiri K, Sawada Y, Kumamoto T, Ueda M, et al. Predictive factors for bile leakage after hepatectomy for hepatic tumors: a retrospective multicenter study with 631 cases at Yokohama Clinical Oncology Group (YCOG). J Hepatobiliary Pancreat Sci. 2017;24:33-41.
- Capussotti L, Ferrero A, Vigano L, Sgotto E, Muratore A, Polastri R. Bile leakage and liver resection: Where is the risk? Arch Surg. 2006;141:690-4.
- Koch M, Garden OJ, Padbury R, Rahbari NN, Adam R, Capussotti L, et al. Bile leakage after hepatobiliary and pancreatic surgery: a definition and grading of severity by the International Study Group of Liver Surgery. Surgery. 2011;149:680–8.
- Lo CM, Fan ST, Liu CL, Lai EC, Wong J. Biliary complications after hepatic resection: risk factors, management, and outcome. Arch Surg. 1998;133:156–61.
- Yamashita Y, Hamatsu T, Rikimaru T, Tanaka S, Shirabe K, Shimada M, et al. Bile leakage after hepatic resection. Ann Surg. 2001;233:45–50.
- Nagano Y, Togo S, Tanaka K, Masui H, Endo I, Sekido H, et al. Risk factors and management of bile leakage after hepatic resection. World J Surg. 2003;27:695–8.

- Honore C, Vibert E, Hoti E, Azoulay D, Adam R, Castaing D. Management of excluded segmental bile duct leakage following liver resection. HPB (Oxford). 2009;11:364–9.
- Patrono D, Tandoi F, Romagnoli R, Salizzoni M. Excluded segmental duct bile leakage: the case for bilio-enteric anastomosis. Updates Surg. 2014;66:115–9.
- 12. Hoekstra LT, van Gulik TM, Gouma DJ, Busch OR. Posthepatectomy bile leakage: how to manage. Dig Surg. 2012;29:48–53.
- Fragulidis G, Marinis A, Polydorou A, Konstantinidis C, Anastasopoulos G, Contis J, et al. Managing injuries of hepatic duct confluence variants after major hepatobiliary surgery: an algorithmic approach. World J Gastroenterol. 2008;14:3049–53.
- Park JH, Oh JH, Yoon Y, Hong SH, Park SJ. Acetic acid sclerotherapy for treatment of a biliary leak from an isolated bile duct after hepatic surgery. J Vasc Interv Radiol. 2005;16:885–8.
- Fukuhisa H, Sakoda M, Hiwatashi K, Iino S, Minami K, Hashiguchi M, et al. Surgical treatment for the excluded bile leakage from Spiegel lobe after right hemihepatectomy: A case report. Int J Surg Case Rep. 2017;39:159–63.
- Sugiura T, Nagino M, Kamiya J, Nishio H, Ebata T, Yokoyama Y, et al. Infraportal bile duct of the caudate lobe: a troublesome anatomic variation in right-sided hepatectomy for perihilar cholangiocarcinoma. Ann Surg. 2007;246:794–8.
- Sadamori H, Yagi T, Matsuda H, Shinoura S, Umeda Y, Fujiwara T. Intractable bile leakage after hepatectomy for hepatocellular carcinoma in 359 recent cases. Dig Surg. 2012;29:149–56.
- Lee Y, Kim SY, Kim KW, Lee SS, Park SH, Byun JH, et al. Contrast-enhanced MR cholangiography with Gd-EOB-DTPA for preoperative biliary mapping: correlation with intraoperative cholangiography. Acta Radiol. 2015;56:773-81.
- Kusano T, Kida H, Nishiwaki Y, Ikematsu Y, Goto K, Kuroda H, et al. Percutaneous sclerotherapy for intractable external biliary fistula after hepatectomy. Int Surg. 2003;88:72–5.
- Reimer P, Schneider G, Schima W. Hepatobiliary contrast agents for contrast-enhanced MRI of the liver: properties, clinical development and applications. Eur Radiol. 2004;14:559–78.
- Fayad LM, Holland GA, Bergin D, Iqbal N, Parker L, Curcillo PG, et al. Functional magnetic resonance cholangiography (fMRC) of the gallbladder and biliary tree with contrast-enhanced magnetic resonance cholangiography. J Magn Reson Imaging. 2003;18:449–60.
- Yun SU, Cheon YK, Shim CS, Lee TY, Yu HM, Chung HA, et al. The outcome of endoscopic management of bile leakage after hepatobiliary surgery. Korean J Intern Med. 2017;32:79–84.
- 23. Kubo N, Harimoto N, Shibuya K, Ishii N, Tsukagoshi M, Igarashi T, et al. Successful treatment of isolated bile leakage after hepatectomy combination therapy with percutaneous transhepatic portal embolization and bile duct ablation with ethanol: a case report. Surg Case Rep. 2018;4:61.
- Alegre Castellanos A, Molina Granados JF, Escribano Fernandez J, Gallardo Muñoz I, Triviño Tarradas FA. Early phase detection of bile leak after hepatobiliary surgery: value of Gd-EOB-DTPA-enhanced MR cholangiography. Abdom Imaging. 2012;37:795–802.
- 25. Cieszanowski A, Stadnik A, Lezak A, Maj E, Zieniewicz K, Rowinska-Berman K, et al. Detection of active bile leak with Gd-EOB-DTPA enhanced MR cholangiography: comparison of 20–25 min delayed and 60–180 min delayed images. Eur J Radiol. 2013;82:2176–82.
- Salvolini L, Urbinati C, Valeri G, Ferrara C, Giovagnoni A. Contrastenhanced MR cholangiography (MRCP) with GD-EOB-DTPA in evaluating biliary complications after surgery. Radiol Med. 2012;117:354–68.
- Sakamoto K, Tamesa T, Yukio T, Tokuhisa Y, Maeda Y, Oka M. Risk factors and managements of bile leakage after hepatectomy. World J Surg. 2016;40:182–9.
- Mutignani M, Forti E, Dokas S, Pugliese F, Fontana P, Tringali A, et al. Endotherapy for bile leaks from isolated ducts after hepatic resection: A long awaited challenge. Dig Liver Dis. 2017;49:893–7.

- Lee SM, Kim HB, Lee IJ. Fluoroscopy-guided transgastric hepaticoantrostomy in a patient with bile leakage associated with biliary obstruction after left hepatic trisectionectomy. J Vasc Interv Radiol. 2015;26:1748–50.
- Kyokane T, Nagino M, Sano T, Nimura Y. Ethanol ablation for segmental bile duct leakage after hepatobiliary resection. Surgery. 2002;131:111–3.
- Sakaguchi T, Shibasaki Y, Morita Y, Oishi K, Suzuki A, Fukumoto K, et al. Postoperative bile leakage managed by interventional intrabiliary ethanol ablation. Hepatogastroenterology. 2011;58:157-60.
- Shimizu T, Yoshida H, Mamada Y, Taniai N, Matsumoto S, Mizuguchi Y, et al. Postoperative bile leakage managed successfully by intrahepatic biliary ablation with ethanol. World J Gastroenterol. 2006;12:3450–2.
- Matsumoto T, Iwaki K, Hagino Y, Kawano K, Kitano S, Tomonari KI, et al. Ethanol injection therapy of an isolated bile duct associated with a biliary-cutaneous fistula. J Gastroenterol Hepatol. 2002;17:807-10.
- Kim HS, Kim TH, Yun EY, Ham HS, Kim HJ, Jeong CY, et al. Segmental bile duct leakage after hepatic resection managed with percutaneous ablation by N-butyl cyanoacrylate. Korean J Hepatobiliary Pancreat Surg. 2012;16:115–9.
- Kataoka M, Ooeda Y, Yoshioka S, Wakatsuki K, Tonooka T, Kawamoto J, et al. Percutaneous transhepatic ethanol ablation for postoperative bile leakage after a hepatectomy. Hepatogastroenterology. 2011;58:988–91.
- Sadakari Y, Miyoshi A, Ohtsuka T, Kohya N, Takahashi T, Matsumoto K, et al. Percutaneous transhepatic portal embolization for persistent bile leakage after hepatic resection: report of a case. Surg Today. 2008;38:668–71.

Hai S, Tanaka H, Takemura S, Sakabe K, Ichikawa T, Kubo S. Portal vein embolization for an intractable bile leakage after hepatectomy. Clin J Gastroenterol. 2012:5:287–91.

AGSurg Annals of Gastroenterological Surgery

- Ikeda A, Noda T, Hatano H, Takata A, Hirota M, Oshima K, et al. [A Case of Intractable Bile Leakage after Surgery for Gallbladder Cancer Successfully Treated by Transcatheter Arterial Embolization]. Gan To Kagaku Ryoho. 2015;42:1734–6.
- Tanaka S, Hirohashi K, Tanaka H, Shuto T, Lee SH, Kubo S, et al. Incidence and management of bile leakage after hepatic resection for malignant hepatic tumors. J Am Coll Surg. 2002;195:484–9.
- 40. Nasr JY, Hashash JG, Orons P, Marsh W, Slivka A. Rendezvous procedure for the treatment of bile leaks and injury following segmental hepatectomy. Dig Liver Dis. 2013;45:433–6.
- Kyokane T, Nagino M, Oda K, Nimura Y. An experimental study of selective intrahepatic biliary ablation with ethanol. J Surg Res. 2001;96:188–96.
- 42. Woo S, Chung JW, Hur S, Joo SM, Kim HC, Jae HJ, et al. Liver abscess after transarterial chemoembolization in patients with bilioenteric anastomosis: frequency and risk factors. AJR Am J Roentgenol. 2013;200:1370-7.

How to cite this article: Kubo N, Shirabe K. Treatment strategy for isolated bile leakage after hepatectomy: Literature review. *Ann Gastroenterol Surg*. 2020;4:47–55. https://doi.org/10.1002/ags3.12303